

JUN 07 2000

NDA 21-015/S-001

Unimed Pharmaceuticals, Inc.
Attention: Judy Athey
Assistant Manager, Regulatory Affairs
Four Parkway North, Suite 200
Deerfield, IL 60015

Dear Ms. Athey:

Please refer to your supplemental new drug application dated April 28, 2000, received May 1, 2000, submitted under section 505(b) of the Federal Food, Drug, and Cosmetic Act for AndroGel™ (testosterone gel).

This supplemental new drug application provides for a change in the patient package insert to remove from the list of patients who should not take AndroGel those individuals with heart, kidney or liver disease.

We have completed the review of this supplemental application and have concluded that adequate information has been presented to demonstrate that the drug product is safe and effective for use as recommended in the agreed upon labeling text. Accordingly, the supplemental application is approved effective on the date of this letter.

The final printed labeling (FPL) must be identical to the submitted draft labeling (patient package insert submitted April 28, 2000).

Please submit 20 copies of the FPL as soon as it is available, in no case more than 30 days after it is printed. Please individually mount ten of the copies on heavy-weight paper or similar material. For administrative purposes, this submission should be designated "FPL for approved supplement NDA 21-015/S-001." Approval of this submission by FDA is not required before the labeling is used.

If a letter communicating important information about this drug product (i.e., a "Dear Health Care Practitioner" letter) is issued to physicians and others responsible for patient care, we request that you submit a copy of the letter to this NDA and a copy to the following address:

MEDWATCH, HF-2
FDA
5600 Fishers Lane
Rockville, MD 20857

We remind you that you must comply with the requirements for an approved NDA set forth under 21 CFR 314.80 and 314.81.

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If you have any questions, call Kim Colangelo, Regulatory Project Manager, at (301) 827-4260. Sincerely,

Susan Allen, MD, MPH
Acting Director
Division of Reproductive and Urologic Drug Products (HFD-580)
Office of Drug Evaluation III
Center for Drug Evaluation and Research

JUN -7 2000



Patient Information and Instructions for Using ANDROGEL™ 1% (testosterone gel) C-III

Read this information carefully before using AndroGel [AN draw jel]. The following information about AndroGel should not take the place of your doctor's orders or recommendations. Your doctor will tell you exactly what dose to take, how to safely take it, and when to take it. Make sure you understand the benefits and risks of AndroGel before you use it. Ask your doctor or pharmacist if you have any other questions about your AndroGel therapy.

What is ANDROGEL?

Your doctor has prescribed this therapy because your body is not making enough testosterone. The medical term for this condition is hypogonadism. Testosterone helps the body produce sperm and the male sexual characteristics. Testosterone is also necessary for normal sexual function and sex drive.

AndroGel is a clear, colorless gel medicine that delivers testosterone into your body through your skin. Once AndroGel is absorbed through your skin, it enters your bloodstream and helps your body reach normal testosterone levels. The type of testosterone delivered into your body by AndroGel is the same as the testosterone produced in your testicles.

Who should not take ANDROGEL?

AndroGel **must not be used by women** or by those individuals with known hypersensitivity to any of its components. Pregnant women should avoid skin contact with AndroGel application sites in men. The active ingredient in AndroGel is testosterone. (See "Inactive Ingredients" at the end of this leaflet for a list of the other ingredients.) Testosterone may cause fetal harm.

You should not use AndroGel if you have any of the following conditions:

- prostate cancer (if your doctor knows for sure or suspects it)
- breast cancer (a rare condition for men)
- difficulty in urinating due to an enlarged prostate

Tell your doctor about other medicines you are taking. AndroGel may affect how these medicines work, and you may need to have your doses adjusted.

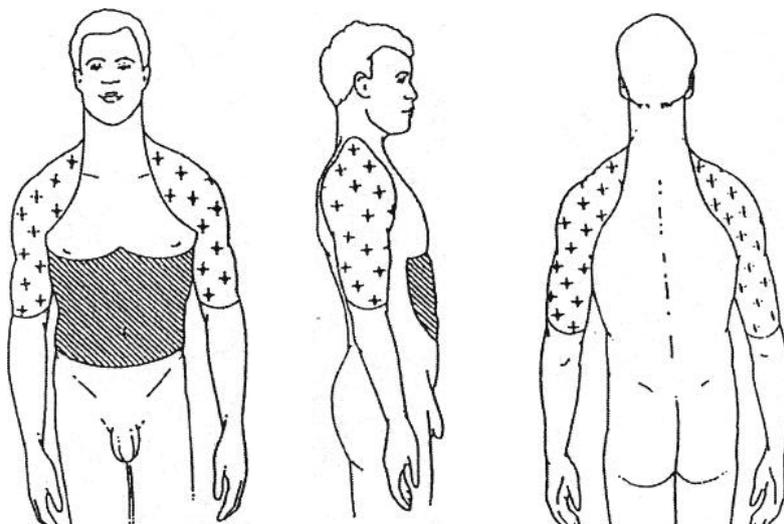
How should I use ANDROGEL?

It is important that you read and follow these directions on how to use AndroGel properly.

1. **Apply AndroGel at the same time (preferably every morning).** You should apply the contents of one (1) packet of gel every morning. If you take a bath or shower in the morning, use AndroGel **after** your bath or shower. Your doctor may tell you to use more than one packet at a time.
2. **Be sure your skin is completely dry.**

3. **Open the packet.** Open one AndroGel aluminum foil packet by folding the top edge and tearing open the packet.
4. **Remove the contents from the packet. Squeeze the entire contents into the palm of your hand. Squeeze from the bottom of the packet toward the top. If you like, you can squeeze out half (1/2) of the contents of the packet at a time and then squeeze out the second half after you have applied the first half, following the directions below.**

Apply gel to starred (upper arm/shoulder) or shaded (abdomen) areas only.



5. **Apply AndroGel only to your abdomen (stomach area), shoulders, or upper arms.** In this way your body will absorb the right amount of testosterone. **Never apply AndroGel to your genitals (penis and scrotum) or to damaged skin.** Apply AndroGel only to healthy, normal skin. Avoid skin with open sores, wounds, or irritation. Use a circular motion to rub the gel for several seconds.
6. **Wash your hands with soap and water right away after application.** Wash your hands right away to reduce the chance that the medicine will spread from your hands to other people.
7. **Let AndroGel dry for a few minutes before you dress.** This prevents your clothing from wiping the gel off your skin. It ensures that your body will absorb the correct amount of testosterone.
8. **Wait 5 to 6 hours before showering or swimming.** To ensure that the greatest amount of AndroGel is absorbed into your system, you should wait 5 to 6 hours after application before showering or swimming. Once in a while, you may shower or swim as soon as 1 hour after applying AndroGel. If done infrequently, this will have little effect on the amount of AndroGel that is absorbed by your body.

9. **Maintain normal activities.** Once your hands are washed and the application site is covered with clothing, there is little risk of transferring testosterone to someone else's skin due to bodily contact. If, however, you expect direct skin contact with someone else, you should wash your application site with soap and water before that encounter. This will reduce the chance that the medicine will transfer to the other person.

What to do if someone else is exposed to AndroGel.

If someone else is exposed to AndroGel either by direct contact with the gel itself or indirectly because of contact with your treated skin, that person should wash the area of contact with soap and water as soon, as possible. The longer the gel is in contact with the skin before washing, the greater is the chance that some testosterone will be absorbed by the other person. This is especially important for women (especially pregnant women) and children. They have naturally low levels of testosterone and could be harmed by it.

What to do if you get AndroGel in your eyes.

If you get AndroGel in your eyes, rinse your eyes right away with warm clean water to flush out any AndroGel. Seek medical attention if needed.

What to do if you miss a dose.

If you miss a dose, do not double your next dose the next day to catch up. If your next dose is less than 12 hours away, it is best just to wait. Do not take the skipped dose. If it is more than 12 hours until your next dose, take the dose you missed. Resume your normal dosing the next day.

What should I avoid while using AndroGel?

It is important that you do not spread the medicine to others, especially women and children. Be sure to wash your hands after applying AndroGel. Do not allow other persons to contact your skin where you have applied AndroGel, especially pregnant or nursing women. **Testosterone may harm the developing baby,**

What are the possible side effects of AndroGel?

AndroGel may cause the following side effects:

- breast development and breast discomfort
- extra fluid in the body. This may cause serious problems for patients with heart, kidney, or liver damage.
- sleep disturbance called "sleep apnea." This is more likely in patients who are overweight or who have lung disease.
- prostate enlargement, sometimes accompanied by difficulty urinating
- emotional problems like depression
- changes in blood levels of cholesterol. This may be monitored and prevented by periodic blood tests.

Tell your doctor if you develop any of the following side effects:

- penis erections that are too frequent or continue too long
- nausea, vomiting, yellow or darker skin (jaundice), or ankle swelling
- breathing problems, including problems breathing while sleeping
- any side effect that concerns you

Tell your doctor if your female partner develops changes in hair distribution, increases in acne, or other signs of masculinity.

Older patients may be at increased risk of developing enlarged prostate or prostate cancer. This also may be monitored by periodic blood tests and prostate exams.

Other Information

Never share your AndroGel with anyone. Every patient is different. Your doctor has prescribed AndroGel specifically for your needs. Use AndroGel only for the condition for which it was prescribed. Medicines are sometimes prescribed for purposes other than those described in a patient information leaflet. If you have any questions or concerns about your AndroGel treatment, ask your health care provider or pharmacist. They can answer your questions and give you the printed information about AndroGel that is written for health professionals.

Inactive Ingredients

Ethanol, purified water, sodium hydroxide, Carbomer 940 and isopropyl myristate.

Store at controlled room temperature 20-25°C (68-77°F) [See USP].

Manufactured by: Laboratoires Besins Iscovesco
Montrouge, France
For Unimed Pharmaceuticals, Inc.
Buffalo Grove, IL 60089

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